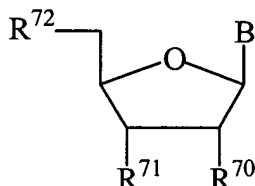


Please cancel claims 1-69 without prejudice.

Please add new claims 70-105 as follows:

-- 70. A labeled nucleoside/tide or nucleoside/tide analog comprising a rhodamine dye conjugated by a linker to a nucleoside/tide or nucleoside/tide analog, wherein:

the rhodamine is a rhodamine-type parent xanthene having attached to the xanthene C9 carbon a phenyl group that is further substituted with an ortho carboxy or ortho sulfonate group or a salt thereof, one to three substituted or unsubstituted aminopyridinium groups and a substituted or unsubstituted alkylthio, or arylthio group; and the nucleoside/tide or nucleoside/tide analog comprises the structure:

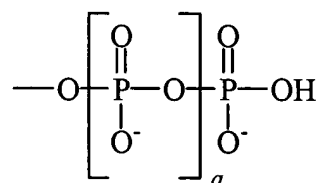


wherein:

B is a nucleobase selected from a purine, a 7-deazapurine, an 8-aza,7-deazapurine, a pyrimidine, a normal nucleobase and a common analog of a normal nucleobase;

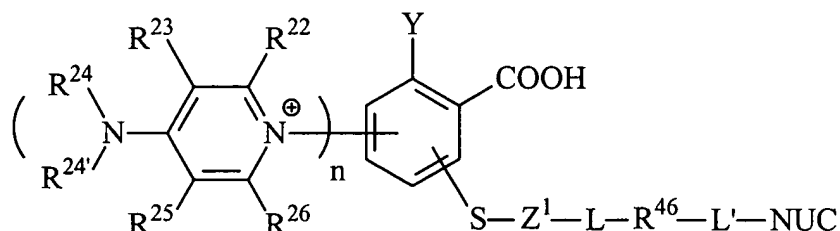
R^{70} and R^{71} , when taken alone, are each independently selected from hydrogen, hydroxyl and a moiety which blocks polymerase-mediated template-directed polymerization, or when taken together form a bond such that the illustrated sugar is 2',3'-didehydroribose; and

R^{72} is selected from hydroxyl, a phosphate ester having the formula:



where a is an integer from 0 to 2, and a phosphate ester analog, or a salt thereof.

71. The labeled nucleoside/tide or nucleoside/tide analog of claim 70 comprising the formula:



wherein:

Y is a rhodamine-type parent xanthene ring attached to the illustrated phenyl group at the xanthene C9 carbon;

R^{22} , R^{23} , R^{25} , and R^{26} are independently selected from hydrogen and (C_1-C_6) alkyl;

R^{24} , when taken alone, is (C_1-C_6) alkyl, or when taken together with $R^{24'}$ is (C_4-C_{10}) alkyldiyl, (C_4-C_6) alkylene, (C_4-C_6) heteroalkyldiyl and (C_4-C_6) heteroalkylene;

$R^{24'}$, when taken alone, is (C_1-C_6) alkyl, or when taken together with R^{24} is (C_4-C_{10}) alkyldiyl, (C_4-C_6) alkylene, (C_4-C_6) heteroalkyldiyl and (C_4-C_6) heteroalkylene;

n is 1, 2, or 3;

S is sulfur;

Z^1 is selected from (C_1-C_{12}) alkyldiyl, (C_1-C_{12}) alkylene independently substituted with one or more of the same or different W^1 groups, (C_5-C_{14}) aryldiyl, and (C_5-C_{14}) aryldiyl independently substituted with one or more of the same or different W^2 groups;

W^1 is selected from $-X$, $-R$, $=O$, $-OR$, $-SR$, $=S$, $-NRR$, $=NR$, $-CX_3$, $-CN$, $-OCN$, $-SCN$, $-NCO$, $-NCS$, $-NO$, $-NO_2$, $=N_2$, $-N_3$, $-S(O)_2O^-$, $-S(O)_2OH$, $-S(O)_2R$, $-C(O)R$, $-C(O)X$, $-C(S)R$, $-C(S)X$, $-C(O)OR$, $-C(O)O^-$, $-C(S)OR$, $-C(O)SR$, $-C(S)SR$, $-C(O)NRR$, $-C(S)NRR$ and $-C(NR)NRR$;

W^2 is selected from $-R$, $-OR$, $-SR$, $-NRR$, $-S(O)_2O^-$, $-S(O)_2OH$, $-S(O)_2R$, $-C(O)R$, $-C(O)X$, $-C(S)R$, $-C(S)X$, $-C(O)OR$, $-C(O)O^-$, $-C(S)OR$, $-C(O)SR$, $-C(S)SR$, $-C(O)NRR$, $-C(S)NRR$ and $-C(NR)NRR$;

L is a selected from a bond, (C₁-C₁₂) alkylidyl, (C₁-C₁₂) substituted alkylidyl, (C₆-C₂₆) arylalkylidyl, -O-, -S-, -NR-, -C(O)O-, -C(O)NR-, -NRS(O)₂-, -NR-NR-, -NRC(O)O-, and -NRC(O)NR-;

R⁴⁶ is selected from -C(O)NR-, -C(O)O-, and -C(O)S-,

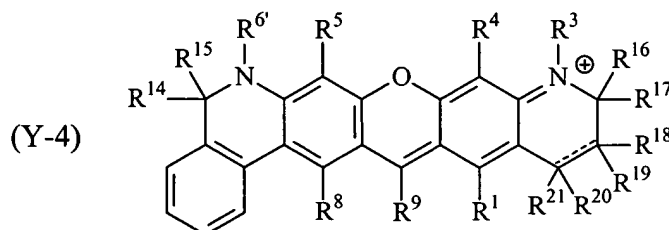
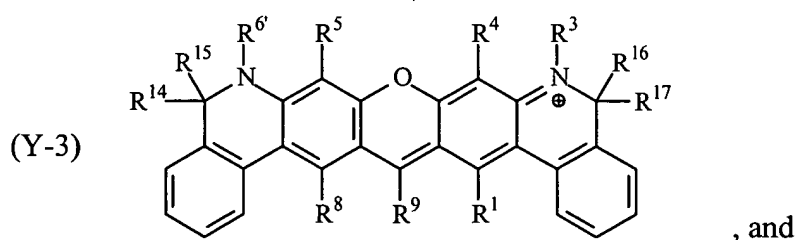
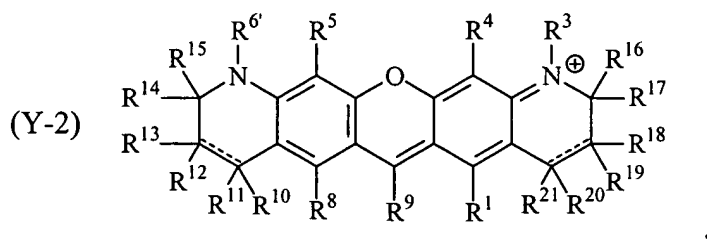
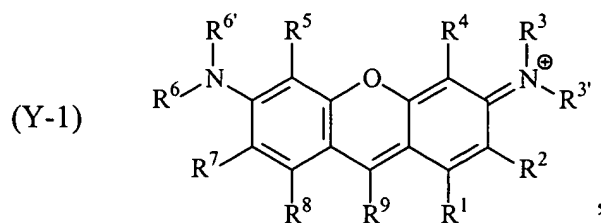
L' is selected from (C₁-C₂₀) alkylidyl, (C₁-C₂₀) heteroalkylidyl, (C₁-C₂₀) alkylene, (C₁-C₂₀) heteroalkylene, (C₆-C₂₆) arylalkylidyl, (C₅-C₂₀) heteroarylalkylidyl, and substituted forms thereof; and

NUC is a nucleoside/tide or nucleoside/tide analog;

each R is independently selected from hydrogen, (C₁-C₆) alkyl, (C₅-C₂₀) aryl, (C₆-C₂₆) arylalkyl, and (C₅-C₂₀) arylaryl; or when two R groups on the same nitrogen atom are taken together, those two R groups are (C₄-C₁₀) alkylidyl or (C₄-C₁₀) alkylene; and

each X is independently a halogen.

72. The labeled nucleoside/tide or nucleoside/tide analog of claim 71 wherein Y comprises the rhodamine-type parent xanthene ring structures:



and a salt thereof, wherein:

R^1 and R^2 when taken alone, are independently hydrogen or (C_1-C_6) alkyl;

R^3 and $R^{3'}$ when taken alone, are independently selected from hydrogen, (C_1-C_6) alkyl, (C_5-C_{14}) aryl and (C_5-C_{14}) arylaryl, or when taken together is (C_4-C_6) alkyldiyl or (C_4-C_6) alkyleno, or when individually taken together with R^2 or R^4 is (C_2-C_6) alkyldiyl or (C_2-C_6) alkyleno;

R^4 , when taken alone, is selected from hydrogen and (C_1-C_6) alkyl, or when taken together with R^3 or $R^{3'}$ is (C_2-C_6) alkyldiyl or (C_2-C_6) alkyleno;

R^5 , when taken alone, is selected from hydrogen and (C_1-C_6) alkyl, or when taken together with R^6 or $R^{6'}$ is (C_2-C_6) alkyldiyl or (C_2-C_6) alkyleno;

R^6 and $R^{6'}$ when taken alone, are selected from hydrogen, (C_1-C_6) alkyl, (C_5-C_{14}) aryl and arylaryl, or when taken together are (C_4-C_6) alkylidyl or alkyleno, or when individually taken together with R^5 or R^7 is (C_2-C_6) alkylidyl or alkyleno;

R^7 , when taken alone, is selected from hydrogen and (C_1-C_6) alkyl, or when taken together with R^6 or $R^{6'}$ is (C_2-C_6) alkylidyl or alkyleno;

R^8 , when taken alone, is selected from hydrogen and (C_1-C_6) alkyl;

R^{10} , R^{11} , R^{12} , R^{13} , R^{14} , R^{15} , R^{16} , R^{17} , R^{18} , R^{19} , R^{20} and R^{21} are each independently selected from hydrogen and (C_1-C_6) alkyl, or

when R^{10} , R^{11} , R^{12} and R^{13} taken together are (C_5-C_{14}) aryleno or (C_5-C_{14}) aryleno substituted with one or more of the same or different (C_1-C_6) alkyl, or

when R^{18} , R^{19} , R^{20} and R^{21} taken together are (C_5-C_{14}) aryleno or aryleno substituted with one or more of the same or different (C_1-C_6) alkyl; and

R^9 is the point of attachment to the xanthene C9 carbon.

73. The labeled nucleoside/tide or nucleoside/tide analog of claim 72 wherein R^2 when taken together with R^3 or $R^{3'}$ is (C_2-C_6) alkylidyl or (C_2-C_6) alkyleno.

74. The labeled nucleoside/tide or nucleoside/tide analog of claim 72 wherein:
an alkylidyl or alkyleno bridge formed by taking R^2 together with R^3 or $R^{3'}$, R^7 together with R^6 or $R^{6'}$, or R^4 together with R^3 or $R^{3'}$, is ethano, propano, 1,1-dimethylethano, 1,1-dimethylpropano or 1,1,3-trimethylpropano;

an aryleno bridge formed by taking R^1 together with R^2 is benzo or naphtho;

an alkylidyl or alkyleno bridge formed by taking R^3 together with $R^{3'}$, or R^6 together with $R^{6'}$, is butano;

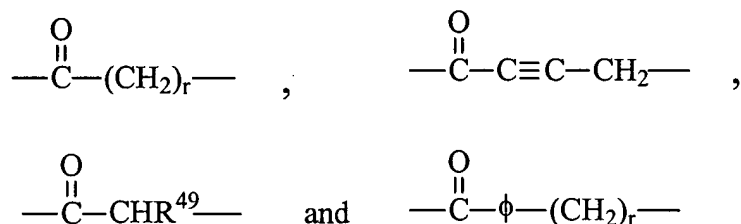
an alkylidyl or alkyleno bridge formed by taking R^5 together with R^6 or $R^{6'}$ is ethano, propano, 1,1-dimethylethano, 1,1-dimethylpropano and 1,1,3-trimethylpropano; and

an aryleno bridge formed by taking R^{10} , R^{11} , R^{12} and R^{13} together, or R^{18} , R^{19} , R^{20} and R^{21} together, is benzo.

75. The labeled nucleoside/tide or nucleoside/tide analog of claim 71 in which Z^1 is phenyldiyl.

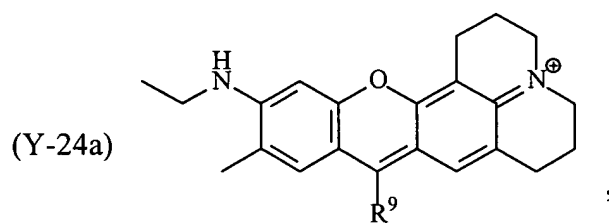
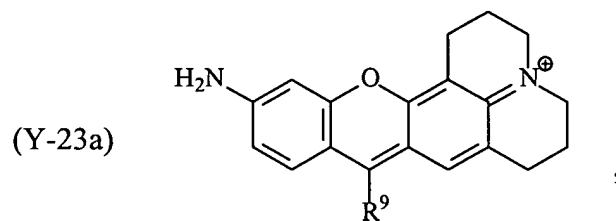
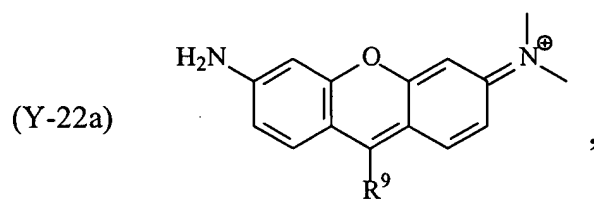
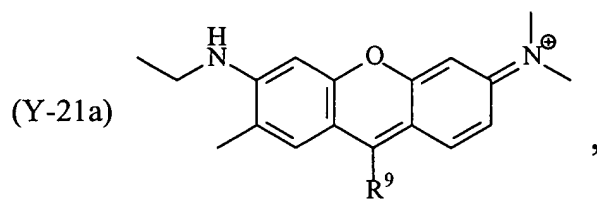
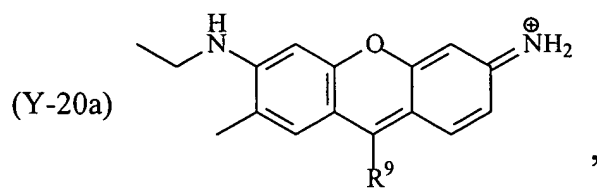
76. The labeled nucleoside/tide or nucleoside/tide analog of Claim 71 in which L' is selected from: $—C\equiv C—CH_2—$ and $—C\equiv C—CH_2—O—CH_2CH_2—$.

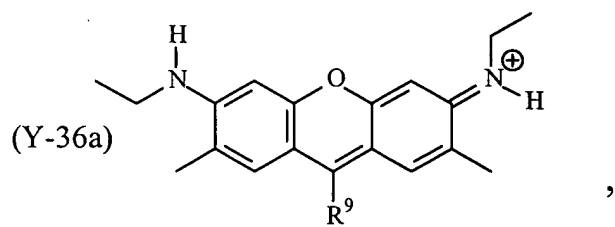
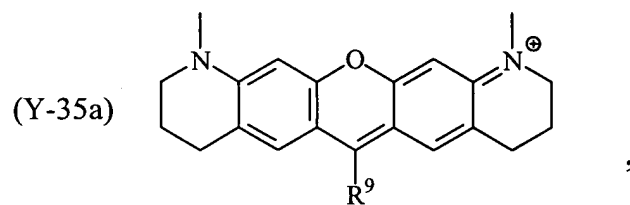
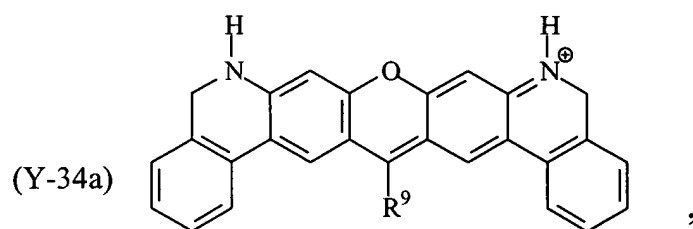
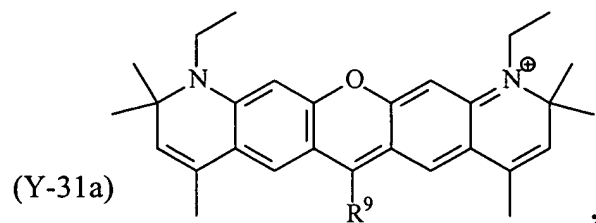
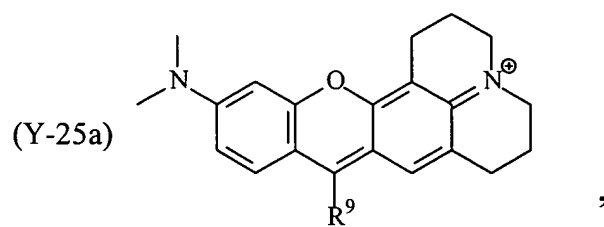
77. The labeled nucleoside/tide or nucleoside/tide analog of Claim 71 in which L' is: $—C\equiv C—CH_2—O—CH_2CH_2—\overset{\overset{R^{47}}{|}}{N}—R^{48}—$ wherein R^{47} is hydrogen or (C_1-C_6) alkyl, and R^{48} is selected from:

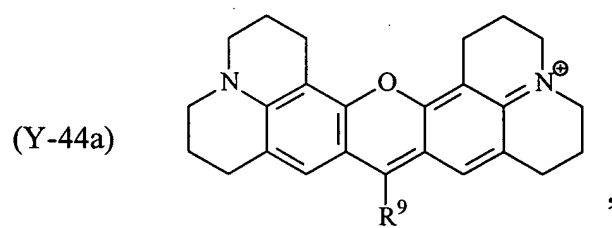
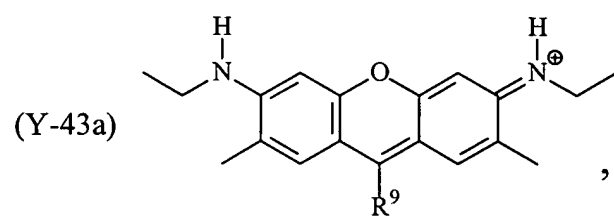
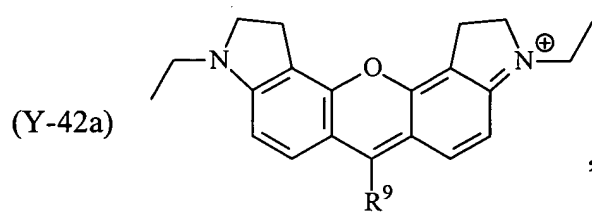
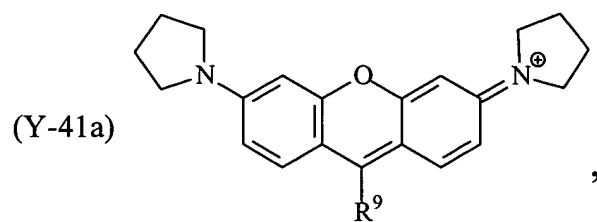
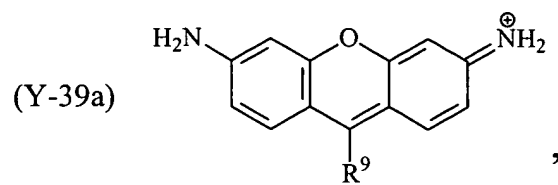


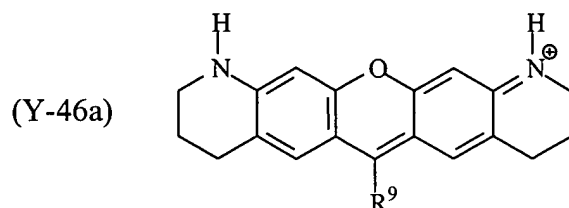
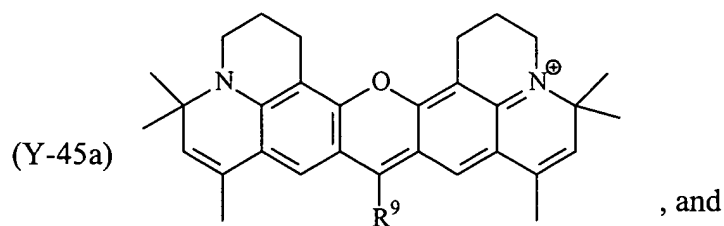
wherein each r is independently an integer from 1 to 6; R^{49} is hydrogen, (C_1-C_6) alkyl, or an amino acid side chain; and ϕ is phenyldiyl or substituted phenyldiyl.

78. The labeled nucleoside/tide or nucleoside/tide analog of claim 71 in which Y is selected from the structures:



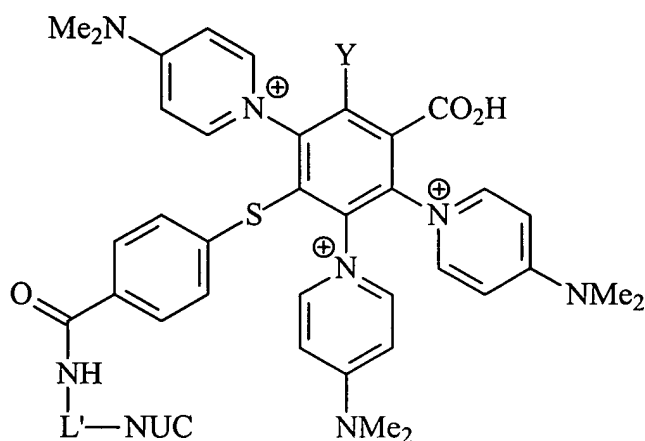






79. The labeled nucleoside/tide or nucleoside/tide analog of claim 71 wherein R^{22} , R^{23} , R^{25} , and R^{26} are each hydrogen.

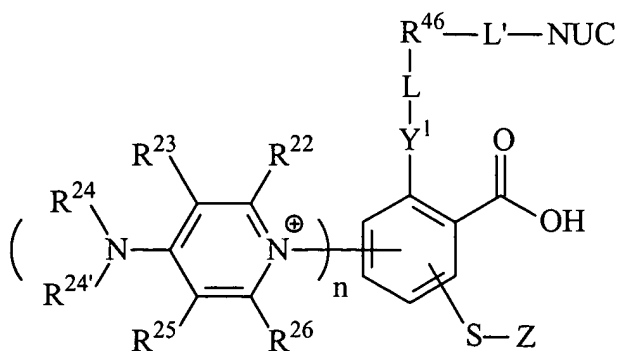
80. The labeled nucleoside/tide or nucleoside/tide analog of claim 71 which comprises the structure:



or a salt thereof.

81. The labeled nucleoside/tide or nucleoside/tide analog of Claim 80 in which L' is selected from: $—C\equiv C—CH_2—$ and $—C\equiv C—CH_2—O—CH_2CH_2—$.

82. The labeled nucleoside/tide or nucleoside/tide analog of claim 70 comprising the formula:



wherein:

Y^1 is a rhodamine-type parent xanthene ring attached to the illustrated phenyl group at the xanthene C9 carbon;

R^{22} , R^{23} , R^{25} , and R^{26} are independently selected from hydrogen and (C₁-C₆) alkyl;

R^{24} , when taken alone, is (C₁-C₆) alkyl, or when taken together with $R^{24'}$ is (C₄-C₁₀) alkyldiyl, (C₄-C₆) alkylene, (C₄-C₆) heteroalkyldiyl or (C₄-C₆) heteroalkylene;

$R^{24'}$, when taken alone, is (C₁-C₆) alkyl, or when taken together with R^{24} is (C₄-C₁₀) alkyldiyl, (C₄-C₆) alkylene, (C₄-C₆) heteroalkyldiyl or (C₄-C₆) heteroalkylene;

n is 1, 2, or 3;

S is sulfur;

Z is (C₁-C₁₂) alkyl, (C₁-C₁₂) alkyl substituted with one or more of the same or different W^1 groups, (C₅-C₂₀) aryl, and (C₅-C₂₀) aryl substituted with one or more of the same or different W^2 groups;

W^1 is selected from -X, -R, =O, -OR, -SR, =S, -NRR, =NR, -CX₃, -CN, -OCN, -SCN, -NCO, -NCS, -NO, -NO₂, =N₂, -N₃, -S(O)₂O⁻, -S(O)₂OH, -S(O)₂R, -C(O)R, -C(O)X, -C(S)R, -C(S)X, -C(O)OR, -C(O)O⁻, -C(S)OR, -C(O)SR, -C(S)SR, -C(O)NRR, -C(S)NRR and -C(NR)NRR;

W^2 is selected from -R, -OR, -SR, -NRR, -S(O)₂O⁻, -S(O)₂OH, -S(O)₂R, -C(O)R, -C(O)X, -C(S)R, -C(S)X, -C(O)OR, -C(O)O⁻, -C(S)OR, -C(O)SR, -C(S)SR, -C(O)NRR, -C(S)NRR and -C(NR)NRR;

L is selected from a bond, (C₁-C₁₂) alkyldiyl, (C₁-C₁₂) substituted alkyldiyl, (C₆-C₂₆) arylalkyldiyl, -O-, -S-, -NR-, -C(O)O-, -C(O)NR-, -NRS(O)₂-, -NR-NR-, -NRC(O)O-, and -NRC(O)NR-;

R^{46} is selected from $-C(O)NR-$, $-C(O)O-$, and $-C(O)S-$,

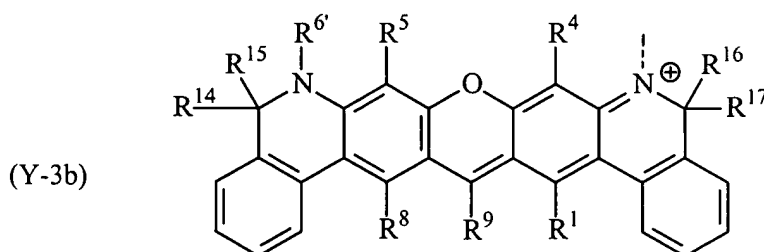
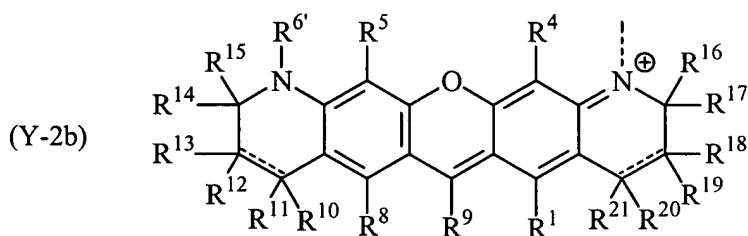
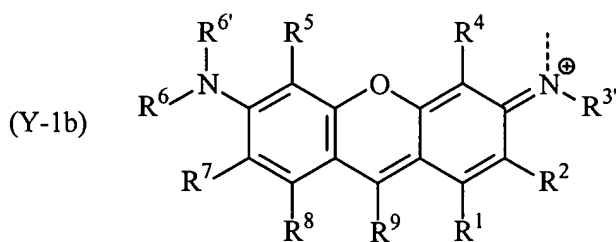
L' is selected from (C_1-C_{20}) alkylidiyl, (C_1-C_{20}) heteroalkylidiyl, (C_1-C_{20}) alkyleno, (C_1-C_{20}) heteroalkyleno, (C_6-C_{26}) arylalkylidiyl, (C_5-C_{20}) heteroarylalkylidiyl, and substituted forms thereof; and

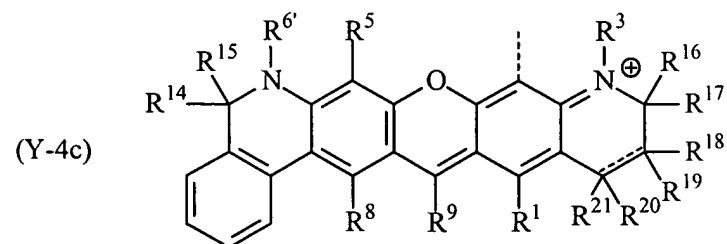
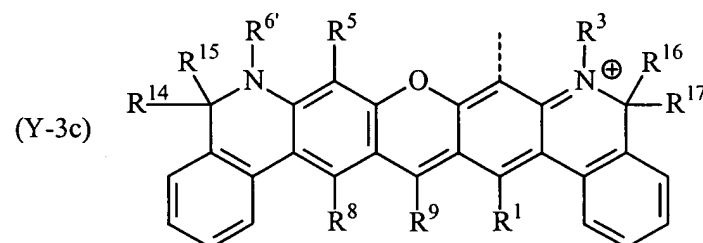
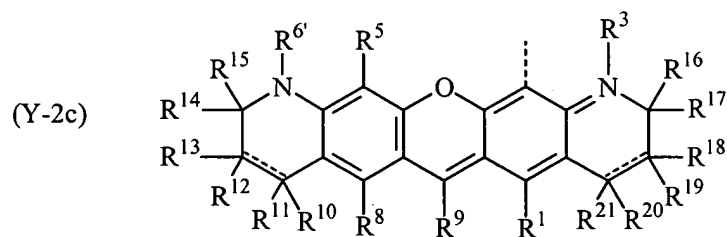
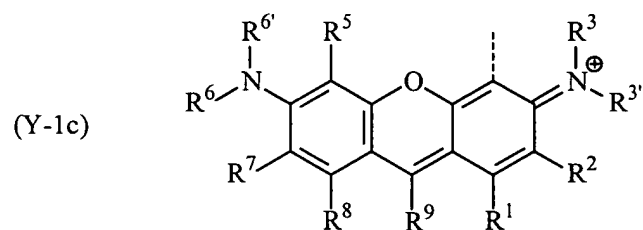
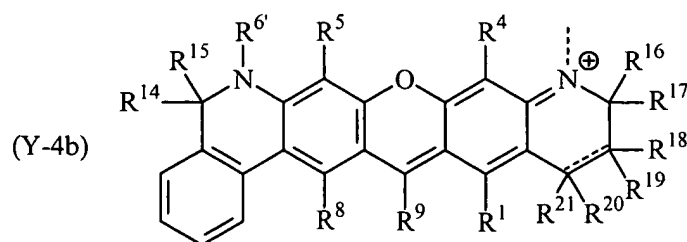
NUC is a nucleoside/tide or nucleoside/tide analog;

each R is independently selected hydrogen, (C_1-C_6) alkyl, (C_5-C_{20}) aryl, (C_6-C_{20}) arylalkyl, and (C_6-C_{20}) arylaryl; or when two R groups on the same nitrogen atom are taken together, those two R groups are (C_4-C_{10}) alkylidiyl or (C_4-C_{10}) alkyleno; and

each X is independently a halogen.

83. The labeled nucleoside/tide or nucleoside/tide analog of Claim 82 in which Y^1 is selected from:





wherein the dashed line at the nitrogen or C4 atom indicates the point of attachment of L.

84. The labeled nucleoside/tide or nucleoside/tide analog of claim 82 wherein:

an alkylidyl or alkylene bridge formed by taking R^2 together with R^3 , R^4 together with R^3 , R^5 together with R^6 , or R^7 together with R^6 , is ethano, propano, 1,1-dimethylethano, 1,1-dimethylpropano or 1,1,3-trimethylpropano; and

an arylene bridge formed by taking R^{10} , R^{11} , R^{12} and R^{13} together or R^{18} , R^{19} , R^{20} and R^{21} together is benzo.

85. The labeled nucleoside/tide or nucleoside/tide analog of claim 82 in which L is selected from phenyldiyl and naphthyldiyl.

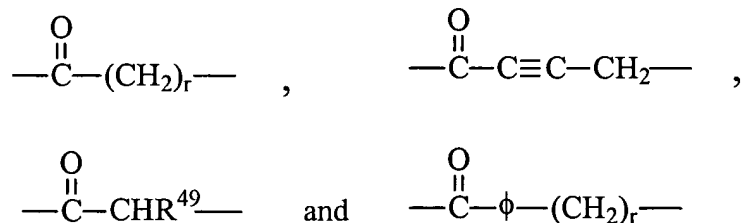
86. The labeled nucleoside/tide or nucleoside/tide analog of claim 82 in which L is $-(CH_2)_i-\phi-$ where i is an integer from 1 to 6 and ϕ is phenyldiyl or naphthyldiyl.

87. The labeled nucleoside/tide or nucleoside/tide analog of claim 82 in which Z is selected from phenyl, benzyl, naphthyl, pyridyl and purinyl.

88. The labeled nucleoside/tide or nucleoside/tide analog of Claim 82 in which L' is selected from: $—C\equiv C—CH_2—$ and $—C\equiv C—CH_2—O—CH_2CH_2—$.

89. The labeled nucleoside/tide or nucleoside/tide analog of Claim 82 in

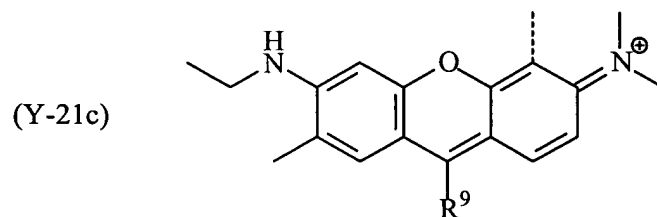
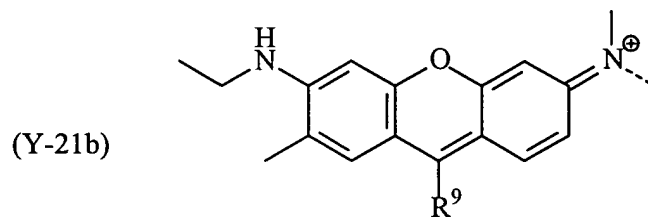
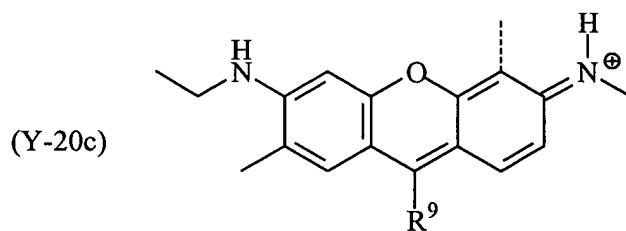
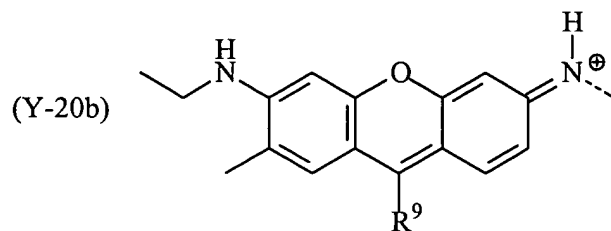
which L' is: $—C\equiv C—CH_2—O—CH_2CH_2—\overset{R^{47}}{\underset{|}{N}}—R^{48}—$ wherein R^{47} is hydrogen or (C_1-C_6) alkyl, and R^{48} is selected from:

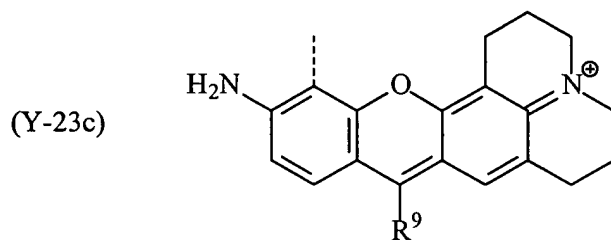
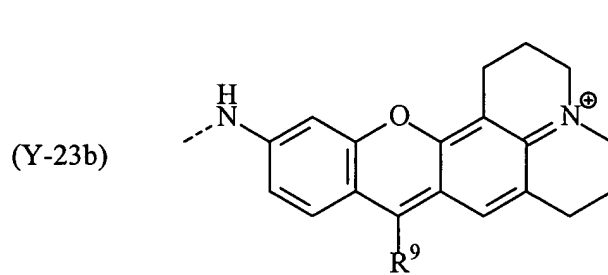
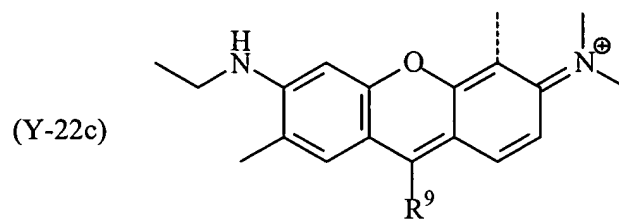
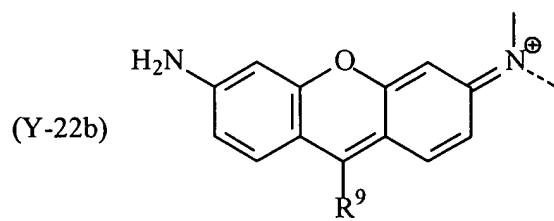


wherein each r is independently an integer from 1 to 6; R^{49} is hydrogen, (C_1-C_6) alkyl, or an amino acid side chain; and ϕ is phenyldiyl or substituted phenyldiyl.

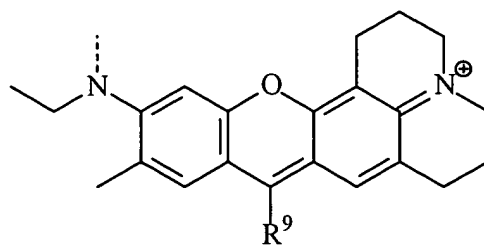
90. The labeled nucleoside/tide or nucleoside/tide analog of claim 82 wherein R^{22} , R^{23} , R^{25} , and R^{26} are each hydrogen.

91. The labeled nucleoside/tide or nucleoside/tide analog of claim 82 in which Y^1 is selected from the group consisting of:

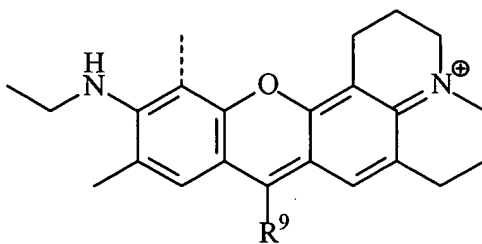




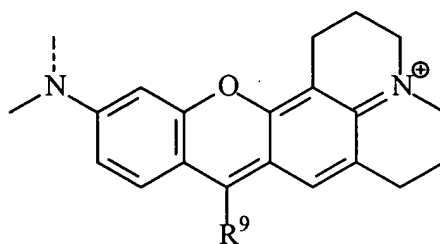
(Y-24b)



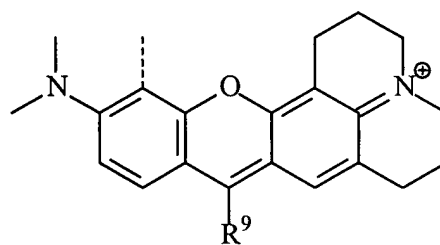
(Y-24c)

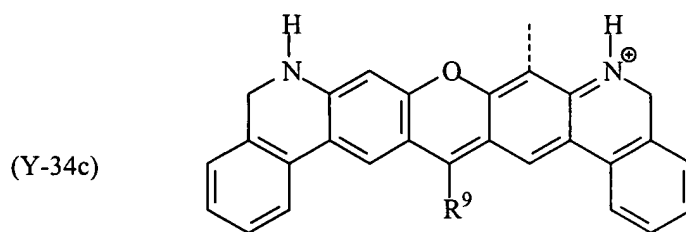
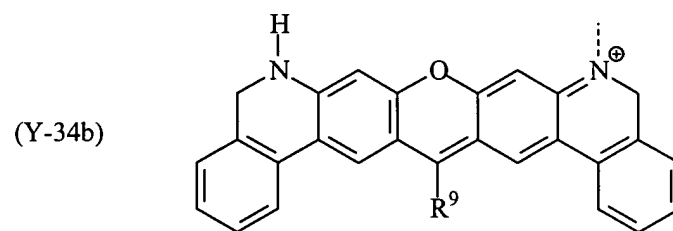
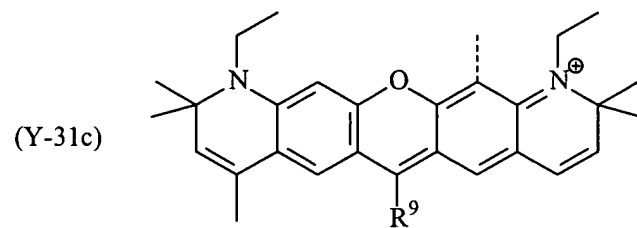
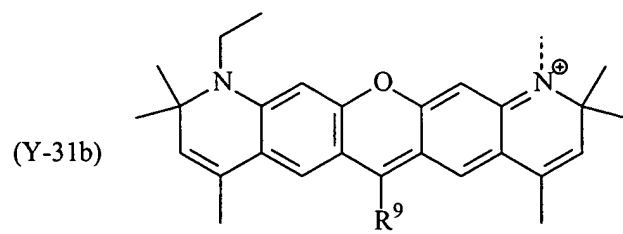


(Y-25b)

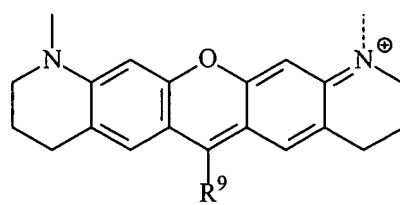


(Y-25c)

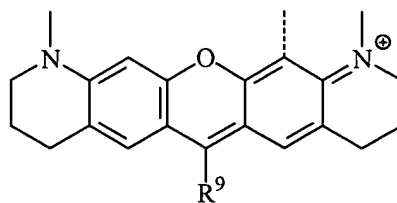




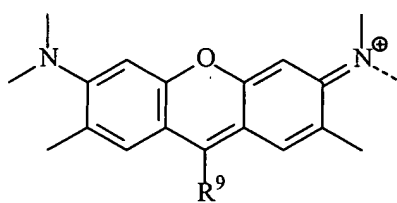
(Y-35b)



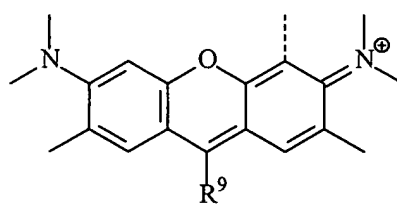
(Y-35c)



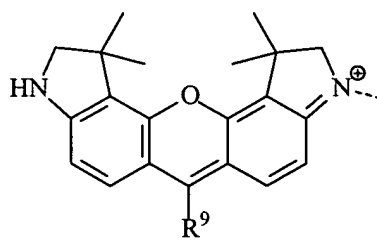
(Y-36b)

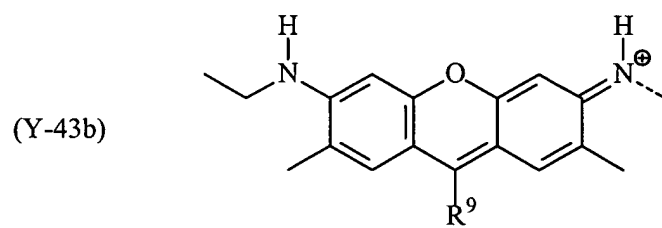
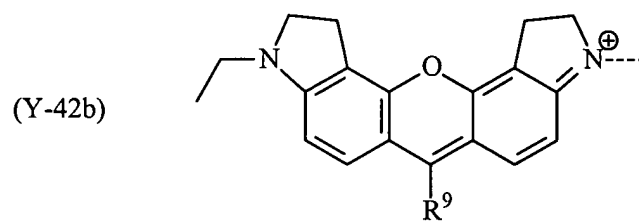
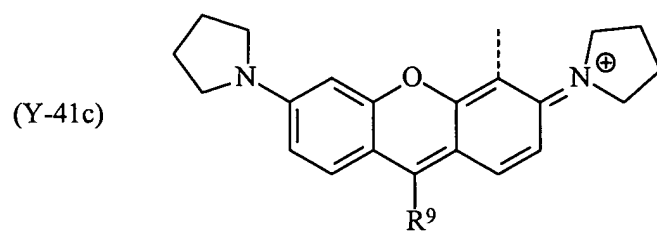
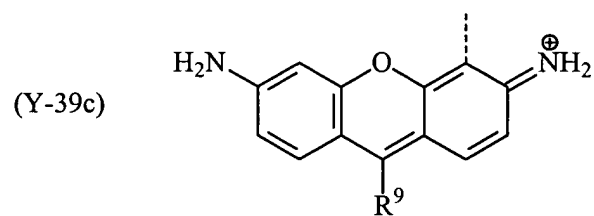
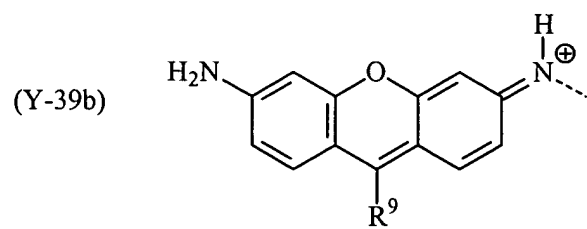


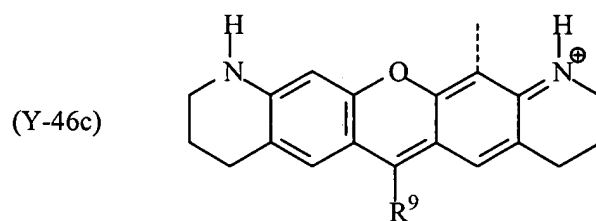
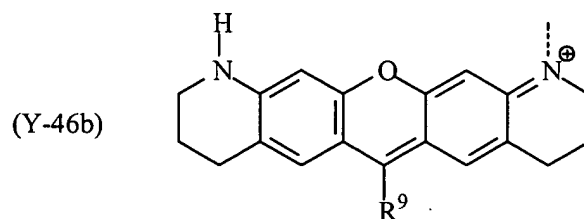
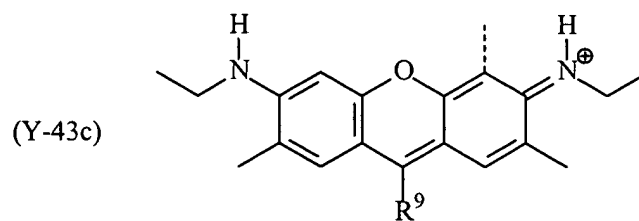
(Y-36c)



(Y-37b)

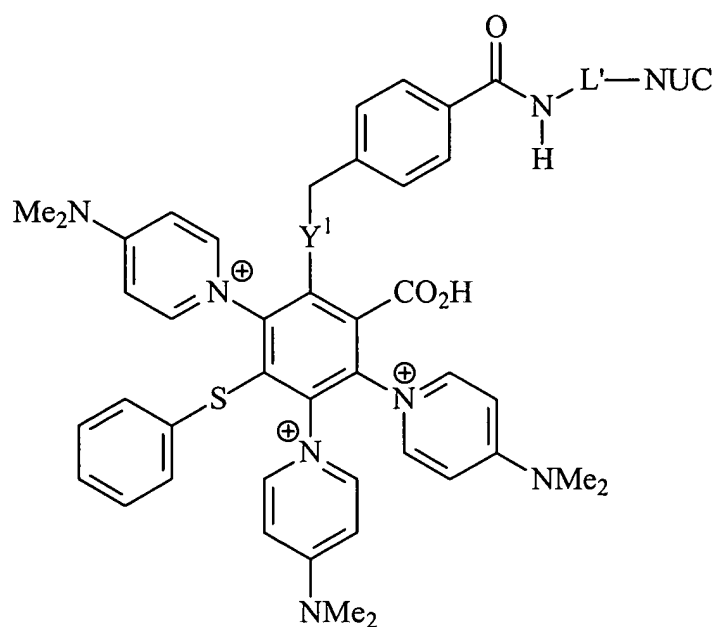






wherein the dash at the nitrogen or C4 atom indicates the point of attachment of L.

92. The labeled nucleoside/tide or nucleoside/tide analog of Claim 82 which has the structure:



93. The labeled nucleoside/tide or nucleoside/tide analog of Claim 92 in which L' is selected from: $\text{—C}\equiv\text{C—CH}_2\text{—}$ and $\text{—C}\equiv\text{C—CH}_2\text{—O—CH}_2\text{CH}_2\text{—}$.

94. The labeled nucleoside/tide or nucleoside/tide analog of Claim 70 further comprising a donor dye or an acceptor dye whereby the rhodamine dye and the donor dye or acceptor dye form an energy-transfer dye pair.

95. The labeled nucleoside/tide or nucleoside/tide analog of Claim 94 wherein the donor dye or acceptor dye is a fluorescein, rhodamine, cyanine, phthalocyanine or squaraine.

96. The labeled nucleoside/tide or nucleoside/tide analog of Claim 94 wherein the donor dye or acceptor dye is 4'-aminomethyl-6-carboxyfluorescein and the 4'-aminomethyl-6-carboxyfluorescein is covalently attached to the rhodamine dye by a linker.

97. The labeled nucleoside/tide or nucleoside/tide analog of Claim 96 wherein the aminomethylfluorescein is further covalently attached by a linker L to the nucleobase B of the nucleoside/tide or nucleoside/tide analog.

98. The labeled nucleoside/tide or nucleoside/tide analog of Claim 70 which is enzymatically incorporatable.

99. The labeled nucleoside/tide or nucleoside/tide analog of Claim 70 which is a terminator.

100. The labeled nucleoside/tide or nucleoside/tide analog of Claim 70 which is enzymatically extendable.

101. The labeled nucleoside/tide or nucleoside/tide analog of Claim 70 wherein R^{71} and R^{70} are hydrogen.

102. The labeled nucleoside/tide or nucleoside/tide analog of Claim 70 wherein R^{71} and R^{70} are hydroxyl.

103. The labeled nucleoside/tide or nucleoside/tide analog of Claim 70 wherein R^{71} is hydroxyl, and R^{70} is hydrogen.